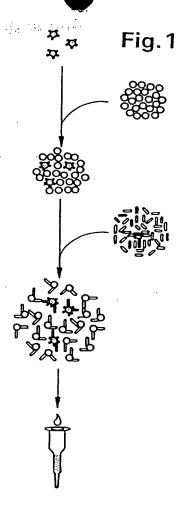
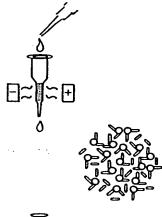
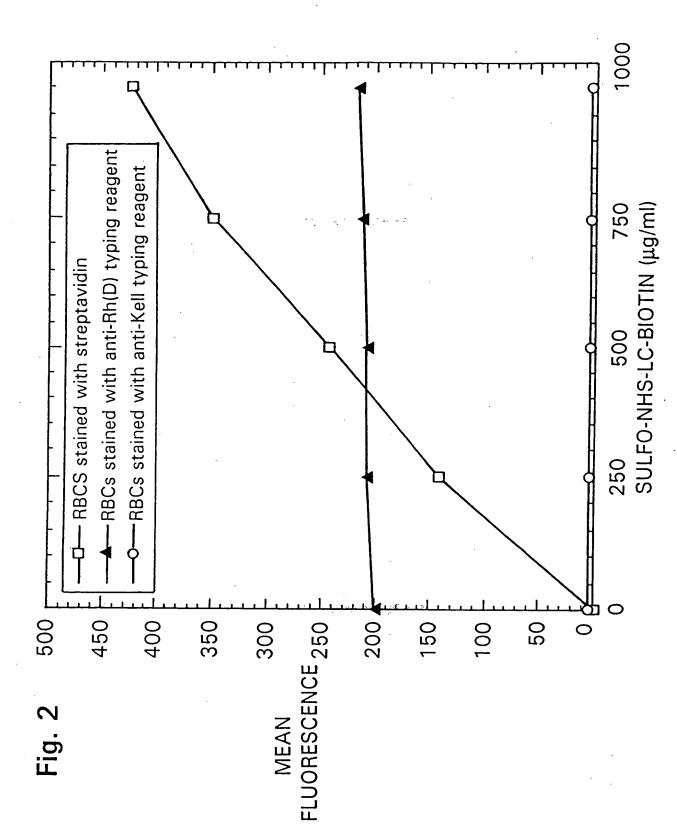
- couple magnetic beads () to antigen-positive cells (②)
- add excess antigen-negative cells (O)
- 3. add phage library containing specificand non-specfic
  - binders
- 4. incubate
- load on column without magnetic field
- 6. place column in magnetic field and wash away antigen-negative cells and non-specific phage
- 7. flush antigenpositive cells
  and bound
  phage from
  column, elute
  bound phage,
  infect bacterial
  culture



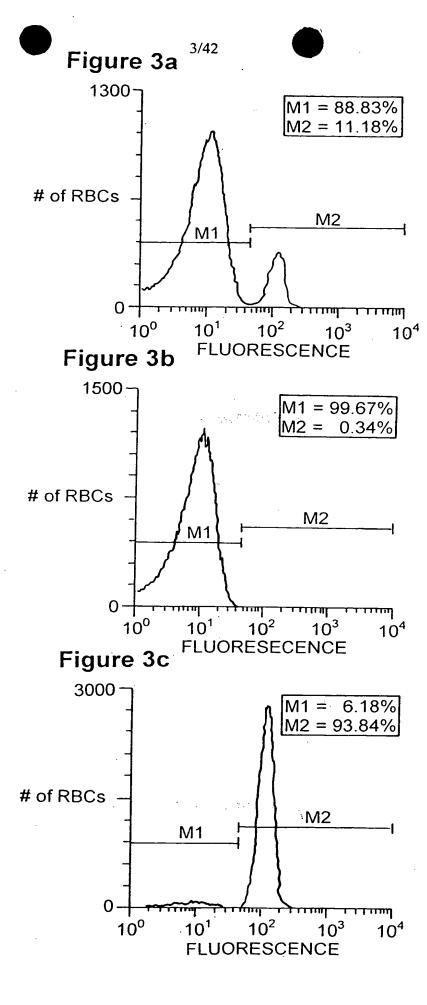




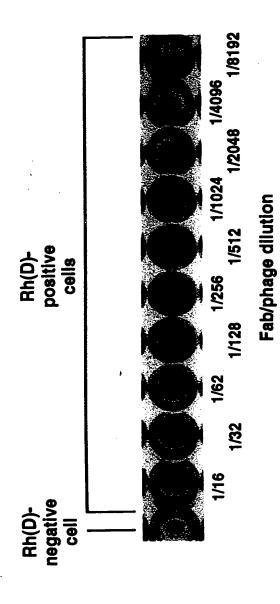




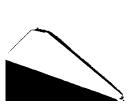








F16.7



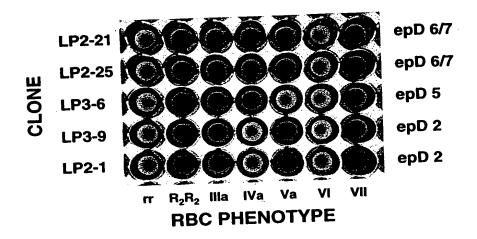
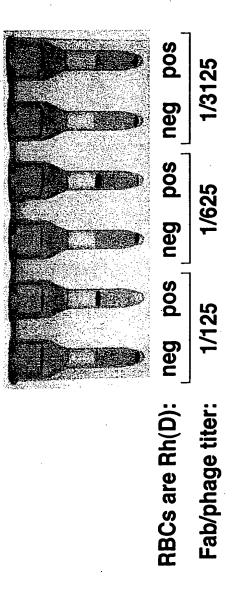
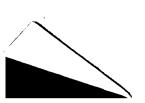


FIG. 5





F16.6



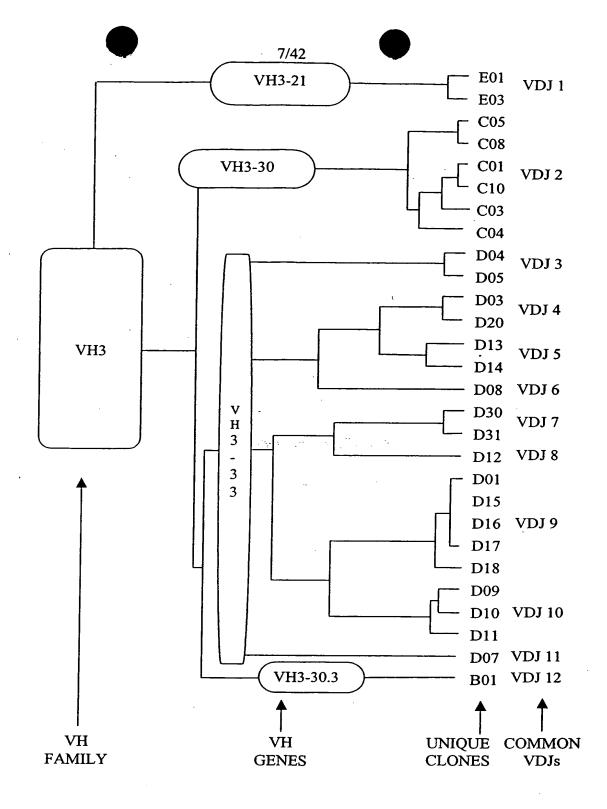


FIG. 7A

CAR DSRYSNFLRWVR-SDGMDV WGQG E01 CAR DSRYSNFLRWVR-SDGMDV WGOG E03 CAN LRGEVTRRASVP----LDI WGQG C05 CAN LRGEVTRRASVP----LDI WGOG C08 CAN LRGEVTRRASVP----FDI WGPG C01 CAN LRGEVTRRASVP----FDI WGPG C10 CAN LRGEVTRRASVP----FDI WGPG C03 CAN LRGEVTRRASIP----FDI WGQG C04 CAR DWR-VRAFS-SGWLSAFDI WGOG D04 CAR DWR-VRAFS-SGWLSAFDI WGQG D05 CAR EEV-VR--GVILWSRKFDY WGQG D03 CAR EEV-VR--GVILWSRKFDY WGQG D20 CAR ENV-ARGGGGVRYKYYFDY WGQG D13 CAR ENV-ARGGGGIRYKYYFDY WGQG D14 CAR DQ---RAAAGIFYYSRMDV WGQG D08 CAR ERN-FR-SGYSRYYYGMDV WGPG D30 CAR ERN-FR-SGYSRYYYGMDV WGPG D31 CAR EAS-ML-RGISRYYYAMDV WGPG D12 CAR ENQ-IK-L-WSRYLYYFDY WGOG D01 CAR ENQ-IK-L-WSRYLYYFDY WGOG D15 CAR ENQ-IK-L-WSRYLYYFDY WGQG D16 CAR ENQ-IK-L-WSRYLYYFDY WGQG D17 CAR ENQ-IK-L-WSRYLYYFDY WGQG D18 CAR EGS-KK-VALSRYYYYMDV WGOG D09 CAR EVS-KK-VALSRYYYYMDV WGOG D10 CAR EVS-KK-LALSRYYYYMDV WGQG D11 CAR ERR-EK--VYILFYSWLDR WGOG D07 CAR GGFYYDSSGYYGLRHYFDS WGQG B01

## FIG. 7B

FIG. 8A

FIG. 8A-3	FIG. 8A-4
FIG. 8A-1	FIG. 8A-2

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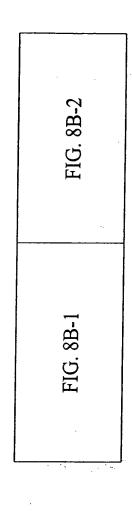
H1	FR1 CDR1 FR2 CDR2	123456789012345678901234567890 1 EVQLVESGGGLVKPGGSLRLSCAASGFTFS S	SE QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VISYDGSNKYYADSVKG  >>>>>>>>>	JH3B QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG >>>>>>> ADDDDDD	JH4B QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG >>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>	JH4B QVQLVESGGGVVQPGRSIRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG >>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>	FIG. 8A-1
		VH D JH  3-21 DA4 JH6B  VDJ1 E01 E03	3-30 DN4 JH3B VDJ2 CA C05 C08 C08 CB C01 C10 C10	3-33 DN1 JF VDJ3 D04 D05	3-33 DXP'1 JF VDJ4 D20 D03	3-33 ?D JE VDJS DA D13 D14	

3-33 DN1 JH6B VDJ6 D08	QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG
3-33 DXP4 JH6B VDJ7 D31 D30	QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG
3-33 DXP'1 JH6B VDJ8 D12	QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG
3-33 DK4 JH4B VDJ9 D15 D16 D01 DB D17 D18	QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG
3-33 DN1 JH6B VDJ10 DC D10 D09 D11	QVQLVESGGGVVQPGRSLRLSCAASGFTFS       SYGMH       WVRQAPGKGLEWVA       VIWYDGSNKYYADSVKG         >>>>>>>>>>>>>>>>>>>>>>>>>>>>>>
3-33 ?D JH5B VDJ11 D07	QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYGMH WVRQAPGKGLEWVA VIWYDGSNKYYADSVKG
3-30.3 PD JH4B VDJ12 B01	QVQLVESGGGVVQPGRSLRLSCAASGFTFS SYAMH WVRQAPGKGLEWVA VISYDGSNKYYADSVKG

	Н3	C	# NUCLEOTIDE
FR3	CDR3	R4	GERMLINE VH
67890123456789012abc345678901234 RFTISRDNAKNSLYLQMNSLRAEDTAVYYCAR	567890abcdefghijk12 ++DYSNY+++YYYYYGMDV DSRYSNFLR-WVRSD	34567890123 WGQGTTVTVSS	<b>ω ω</b>
RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAK  . K T P	+	WGQGTMVTVSS	
RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR ************************************	LKGEVIKKASVP* ++++GYSSSWY++DAFDI DWRVRAFSSGWLS DWRVRAFSSGWLS	WGQGTMVTVSS	
RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR	ITMVRGVII+++++YFDY EEVVRGVILWSRK EEVVRGVILWSRK	WGQGTLVTVSS	S · ·
RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR	+++++++++++++PDY ENVARGGGG?RYKY ENVARGGGURYKY ENVARGGGIRYKY	WGQGTLVTVSS	

	15	111	14	6 0 0 0 0 0 7 T T T T T T T T T T T T T T	1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	23	ω
	WGQ11V1V2W	WGQGTTVTVSSPP	WGQGTTVTVSS	WGQGTLVTVSS	WGQGTTVTVSS	WGQGTLVTVSS	WGQGTLVTVSS
	TGIRAAG-++11111GNDV DQRAAAGIF**SR	YYDFWSGYYTYYYYYGMDV ERNFRSGYSR	+ITMVRGVIIYYYYYGMDV EASMLRGISRA	++WIQLWL++++++YFDY ENQIKLWSRYLY ENQIKLWSRYLY ENQIKLWSRYLY ENQIKLWSRYLY ENQIKLWSRYLY	+++GYSSSWYYYYYGMDV WGQGTTVTVSS EVSKK?ALSRY** EVSKKVALSR*Y** EGSKKVALSR*Y** EGSKKVALSR*Y**	++++++++++++NWFDP WGQGTLVTVSS ERREKVYILFYS.L.R	+++++++++++++YFDY WGQGTLVTVSS GGFYYDSSGYYGLRHS
RFTISRDNSKNTLYLOMNSLRAFINTAVVVCAR		RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR 3	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR + ***********************************	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR +V	RFTISRDNSKNTLYLQMNSLRAEDTAVYYCAR +AVK*FTI E	RFTISRDNSKNTLYLOMNSLRAEDTAVYYCAR +

FIG. 8B

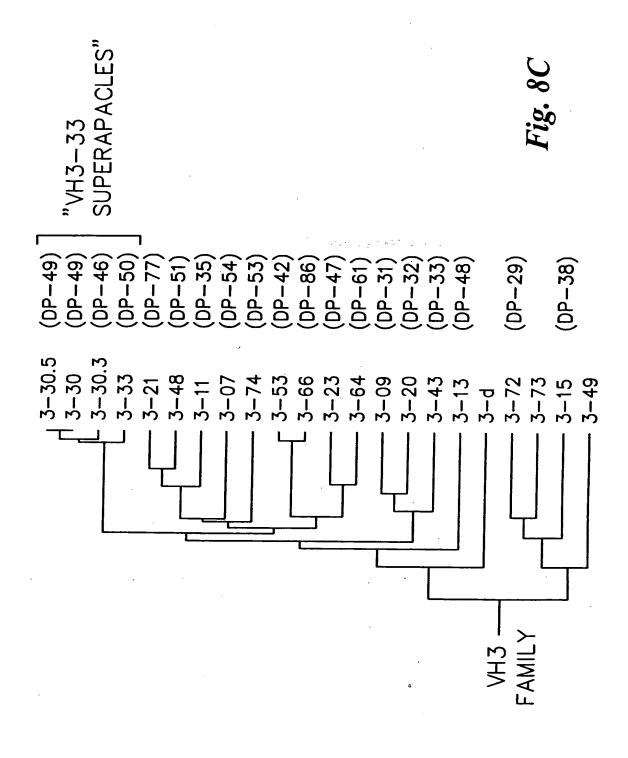


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## S--YGMH WVRQAPGKGLEWVA 1AB2345 . - - . G. . - -QVQLVESGGGVVQPGRSLRLSCAASGFTFS HOMOLOGY TO CON. 0000 0000 0000 3-30.3 3-21 3-33 K

CHOTHIA	. 1-3	1-3	1-3	1-3
5	Ss58.Y1AS	Ж	······································	VISYDGSNKYYADSVKG RFTISRDNSKNTLYLOMNSLRAEDTAVYYCAR
56 012ABC3456789012345	SS. Y.L		· · · · · · · · · · · · · · · · · · ·	VISYDGSNKYYADSVKG

## FIG. 8B-2



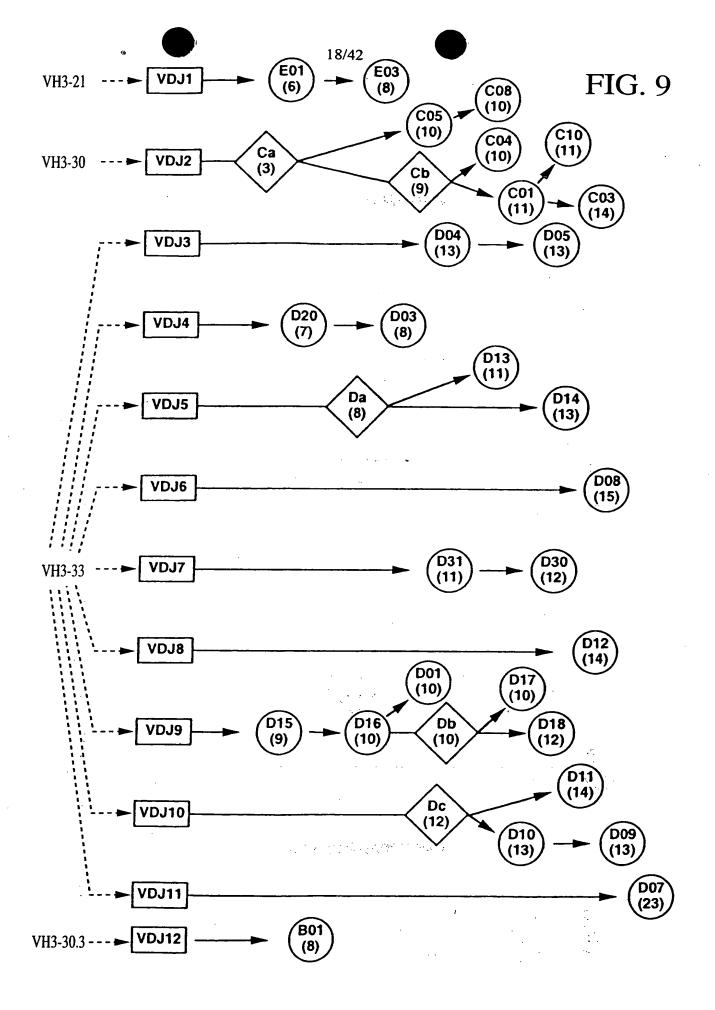


FIG. 10A

FIG. 10A-3	FIG. 10A-4
FIG. 10A-1	FIG. 10A-2

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L2  CDR2	5 9 0123456 Y AASSLQS F GV.KN	Y AASSLOS	Y AASSLQS	Y AASSLQS
FR2	567890123456789 WYQQKPGKAPKLLIY HF.	WYQQKPGKAPKLLIYH.E.	WYQQKPGKAPKLLIY	WYQQKPGKAPKLLIY
L1 CDR1	45678901abcdef234 RASQSISSYLN N.RRS .N.N.RRS	RASQSISSYLNNT.RST.RS	RASQSISSYLN	RASQSISSYLN
FR1	12345678901234567890123 DIQMTQSPSSLSASVGDRVTITC >>>>>>>> >>>>>>>>>>>>>>>>>>>>>>>>>>	DIQMTQSPSSLSASVGDRVTITC >>>>>>>	DIQMTQSPSSLSASVGDRVTITC >>>>>>> >>>>>>	DIQMTQSPSSLSASVGDRVTITC >>>>>>
	<b>J</b> K1	JK2	JK3	JK4
	DPK9 105 104 115 116	DPK9 112 113 108 108 111	DPK9 <b>101</b> <b>103</b>	DPK9 <b>I07</b>

## FIG 10A-1

DPK9 <b>I06</b>	JK5	DIQMTQSPSSLSASVGDRVTITC RASQSISSYLN WYQQKPGKAPKLLIY AASSLQS
DPK8 <b>H01</b>	JK3	DIQLTQSPSFLSASVGDRVTITC RASQGISSYLA WYQQKPGKAPKLLIY AASTLQS
A30 <b>F01</b>	JK1	DIQMTQSPSSLSASVGDRVTITC RASQGIRNDLG WYQQKPGKAPKRLIY AASSLQS
DPK15 <b>G01</b>	JK4	DIVMTQSPLSLPVTPGEPASISC RSSQSLLHSNGYN-YLD WYLQKPGQSPQLLIY LGSNRAS

## FIG. 10A-2

<pre># nucleotide differences from germline Vk</pre>	111 20 44 99	17 173 173 173	13
# diff FR4 g	10345678 89012345678 FGQGTKVEIK	FGQGTKLEIK	FGPGTKVDIKEML. FGGGTKVEIK
L3 CDR3	9012345a67 QQSYSTP+WT *SN.* .TSA.*	QQSYSTP+YT	QQSYSTP+FTP. QQSYSTP+LTR.
FR3	78901234567890123456789012345678 GVPSRFSGSGSGTDFTLTISSLQPEDFATYYC	GVPSRFSGSGTDFTLTISSLQPEDFATYYC  L  P  S  S  S  S  S  S  S  S  S  S  S  S	GVPSRFSGSGSGTDFTLTISSLQPEDFATYYCT

FIG. 10A-3

## ω α FGQGTRLEIK FGPGTKVDIK FGQGTKVEIK FGGGTKVEIK QQSYSTP+IT GVPSRFSGSGGTEFTLTISSLQPEDFATYYC QQLNSYP+FT GVPSRFSGSGSGTEFTLTISSLQPEDFATYYC LQHNSYP+WT GVPDRFSGSGSGTDFTLKISRVEAEDVGVYYC MQALQTP+LT ...N.\*P. GVPSRFSGSGSGTDFTLTISSLQPEDFATYYC

## IG. 10A-4

FIG. 10B-2	
FIG. 10B-1	·

V<sub>K</sub>

DIQMTQSPSSLSASVGDRVTITC RASQSISS-

GENE
DPK9
DPK8
A30
DPK15

# WYQQKPGKAPKLLIY AASSLQS GVPSRFSGSGSGTDFTLTISSLQPEDFATYYC QQSYSTP .LN.Y. L.HN.Y. 표.

## FIG. 10B-2

....K..RVEA..VGV... M.ALQ..

..L....QS.Q.... LG.NRA.

FIG. 11A-3	FIG. 11A-4
FIG. 11A-1	FIG. 11A-2

FR2	567890123456789	WFQQKPGQAPRALIY	WYQQHPGKAPKLMIY *TH	WYQQHPGKAPKLMIY Y*I	WYQQLPGTAPKLLIY *H	WYQQLPGTAPKLLIY *
CDR1	45678901abc234	ASSTGAVTSGYYPNR.F	TGTSSDVGGYNYVS	TGTSSDVGSYNLVS	TGSSSNIGAGYDVH	SGSSSNIGNNY-VS
FR1	1234567891234567890123	QTVVTQEPSLTVSPGGTVTLTC >>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>	QSALTQPPSASGSPGQSVTISC>>>>>>>	QSALTQPASVSGSPGQSITISC	QSVVTQPPSVSGAPGQRVTISC >>>>>>>> >>>>>>>	QSVLTQPPSVSAAPGQKVTISC >>>>>>> >>>>>
•	J <b>À</b>	JL2Vasicek	JL2Vasicek	JL2Vasicek	JL2Vasicek	JL2Vasicek
	۸۸	7a.2.3/DPL18 K01 K02 K03	2c.118D9+ <b>R01</b>	DPL10/lv2066 <b>S01</b>	DPL7/VL1.2 003 002 001	1b.366F5/DPL5 NO2 NO1

# FIG. 11A-1

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<pre>1g.400B5/DPL3 JL2Vasicek</pre>	JL2Vasicek	OSVLTQPPSASGTPGQRVTISC SGSSSNIGSNY-VY WYQQLPGTAPKLLIY >>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>
1c.10.2/DPL2 LO5 LO3 LO4 LO1	JL2Vasicek	OSVLTQPPSASGTPGQRVTISC SGSSSNIGSNT-VN WYQQLPGTAPKLLIY >>>>>>>>>>> >>>>>>>>>>>>>>>>>>>>>>>
DPL16/VL3.1 <b>JO2</b> <b>JO5</b> <b>JO5</b>	JL2Vasicek	SSELTQDPAVSVALGQTVRITC QGDSLRSYYAS WYQQKPGQAPVLVIY >>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>>
3p.81A4+ <b>P01</b>	JL2Vasicek	SYELTQPPSVSVSPGQTARITC SGDALPKKYAY WYQQKSGQAPVLVIY
4b.68B6 <b>201</b>	JL2Vasicek	QLVLTQSPSASASLGASVKLTC TLSSGHSSYAIA WHQQQPEKGPRYLMK

# FIG. 11A-2

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CDR2	FR3	CDR3	ge FR4	germline VA	
5	78	.99 1012345abcdef67 LLYYGGAQ+++++vV SW*	.10 8901234567 FGGGTKLTVL		_
	*	F.AWA		12	•
EVSKRPS GVPDRFSGSKSG .GT	GVPDRFSGSKSGNTASLTVSGLQAEDEADYYC	SSYAGSNNF++++VV F.*NSVI	FGGGTKLTVL	17	30/42
EGSKRPS GVSNRFSGSKSGSRGNSNRPS GVPDRFSGSKSG	GVSNRFSGSKSGNTASLTISGLQAEDEADYYCSRH. GVPDRFSGSKSGTSASLAITGLQAEDEADYYC	CSYAGSSTF++++VVIRI QSYDSSLSG+++VV	FGGGTKLTVL  FGGGTKLTVL	10	
				က	
	· · · · · · · · · · · · · · · · · · ·			10 13	
DNNKRPS GIPDRFSGSKSGTXR*	SGTSATLGITGLQTGDEADYYC	GTWDSSLSA++++VVGRVRRM ADNGR*	FGGGTKLTVL	1.5	
	FIG. 11	11A-3			

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3 6 23	8 18 14 18	25 26 18 21	41	38	
RNNQRPS GVPDRFSGSKSGTSASLAISGLRSEDEADYYC AAWDDSLSG++++VV FGGGTKLTVL	SNNQRPS GVPDRFSGSKSGTSASLAISGLQSEDEADYYC AAWDDSLNG++++VV FGGGTKLTVLK *TR*G *T* TG *T* TG *T*  TG *T*  TG *T*  T	GKNNRPS GIPDRFSGSSSGNTASLTITGAQAEDEADYYC NSRDSSGNH+++++VV FGGGTKLTVL  .R	EDSKRPS GIPERFSGSSSGTMATLTISGAQVEDEADYYC YSTDSSGNH+++++VV FGGGTKLTVL	LNSDGSHSKGD GIPDRFSGSSSGAERYLTISSLQSEDEADYYC QTWGTGI++++++VV FGGGTKLTVL VTNR.IA*SGG.*MH**	

TG. 11B

FIG. 11B-2
FIG. 11B-1

## 3

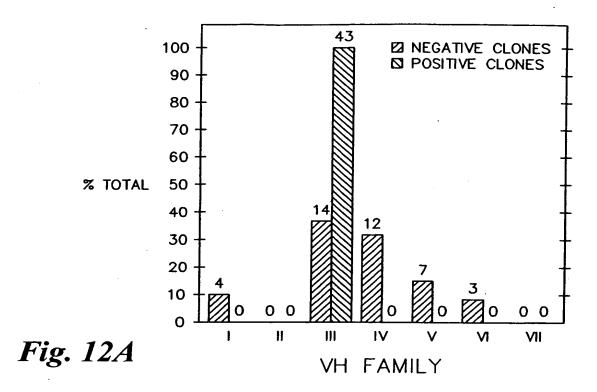
> нн	
PL18 + 2066 .2 /DPL5 /DPL3 DPL2 3.1	ΔΙ
GENE 7a.2.3/DPL18 2c.118D9+ DPL10/1v2066 DPL7/VL1.2 1b.366F5/DPL5 1g.400B5/DPL3 1c.10.2/DPL2 DPL16/VL3.1 3p.81A4+	4D.68B6

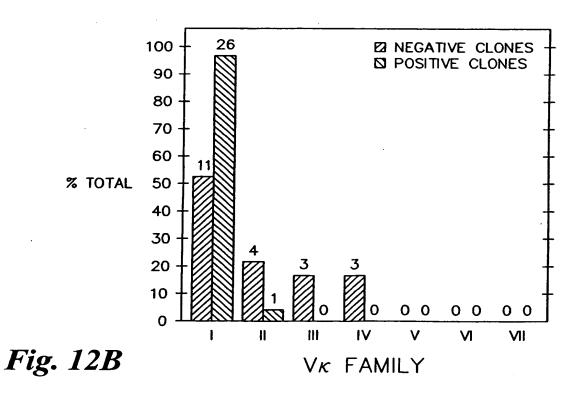
TGTSSDVGGYNYVS TGTSSDVGSYNLVS ASSTGAVTSGYYPN TGSSSNIGAGYDVH SGSSSNIGNNY-VS QGDSLR---SYYAS TLSSG--HSSYAIA SGSSSNIGSNT-VN SGDALP - - - KKYAY SGSSSNIGSNY-VY QTVVTQEPSLTVSPGGTVTLTC QSVVTQPPSVSGAPGQRVTISC QLVLTQSPSASASLGASVKLTC QSALTQPPSASGSPGQSVTISC QSALTQPASVSGSPGQSITISC QSVLTQPPSVSAAPGQKVTISC QSVLTQPPSASGTPGQRVTISC SSELTQDPAVSVALGQTVRITC SYELTQPPSVSVSPGQTARITC QSVLTQPPSASGTPGQRVTISC

## FIG. 11B-1

SSYAGSNNF OSYDSSLSG AAWDDSLNG CSYAGSSTF GTWDSSLSA AAWDDSLSG NSRDSSGNH YSTDSSGNH OTWGTGI GIPDRFSGSSSG--AERYLTISSLOSEDEADYYC GIPERFSGSSSG--TMATLTISGAQVEDEADYYC GVSNRFSGSKSG--NTASLTISGLQAEDEADYYC GVPDRFSGSKSG--TSASLAITGLQAEDEADYYC GVPDRFSGSKSG--TSASLAISGLRSEDEADYYC GVPDRFSGSKSG--TSASLAISGLQSEDEADYYC GIPDRFSGSSG--NTASLTITGAQAEDEADYYC WTPARFSGSLLG--GKAALTLSGVQPEDEAEYYC GIPDRFSGSKSG--TSATLGITGLQTGDEADYYC GVPDRFSGSKSG--NTASLTVSGLQAEDEADYYC --SNKHS --SKRPS SKRPS --SNRPS --NKRPS NORPS --NQRPS GK-----NNRPS ED----SKRPS LNS-DGSHSKGD ---NO RN---EV--GN--SN-. WFQQKPGQAPRALIY WHQQQPEKGPRYLMK WYQQKSGQAPVLVIY WYQQHPGKAPKLMIY WYQQHPGKAPKLMIY WYQQKPGQAPVLVIY WYQQLPGTAPKLLIY WYQQLPGTAPKLLIY WYQQLPGTAPKLLIY WYQQLPGTAPKLLIY

IG. 11B-2





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Fig. 12C

CLONE (HC/LC)	Rh(D)VARIANT CATEGORY	ASSIGNED EPITOPE
	IIIc IVa IVb Va VI VII	
E1/L4	600005	epD1
E1/M2		epD2
E1/M3	00000	epD3
D20/K3	00000	epD6/7
D7/J4	00000	"epDX"

VDJ10 VDJ11 VDJ12 
 VDJ1
 VDJ2
 VDJ3
 VDJ4
 VDJ5
 VDJ5
 VDJ7
 VDJ8
 VDJ9
 VDJ10
 VDJ11
 VDJ11</t 7, HEAVY CHAIN M2 M3 03 01 N2 N1 3 47 3 S الح 25 **R1** 7 λ LIGHT CHAIN

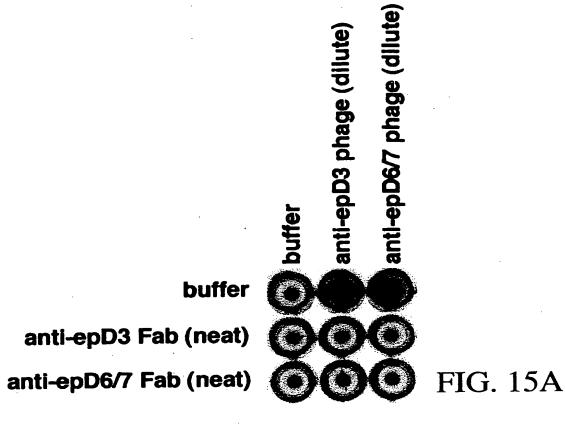
Fig. 14A

 
 VDJ1
 VDJ2
 VDJ3
 VDJ4
 VDJ5
 VDJ5
 VDJ7
 VDJ9
 VDJ9
 VDJ1
 <th 7, HEAVY CHAIN 115 118 112 110 113 ಬ 4 <u></u> 2  $\overline{\omega}$ 9 되도 <u>თ</u> 7  $\overline{c}$ K LIGHT CHAIN

Fig. 14B

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in the same with the same



buffer

anti-epD6/7 Fab (neat)

anti-B Fab (neat)

buffer anti-epD6/7 phage (dilute)

FIG. 15B

diluted diluted	) neat diluted diluted	anti-A anti-A anti-k anti-k
anti-epD3 Fab (γ <sub>η</sub> λ)	anti-epD6/7 Fab (ૠ<)	developed with:

FIG. 15C

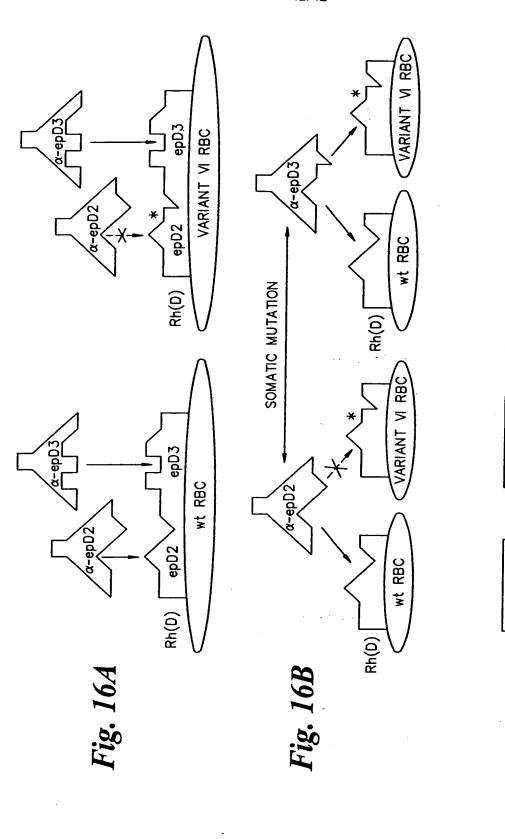


Fig. 16C

PRECURSOR
—— ANTI-epD3

PRECURSOR
—— ANTI-epD3

ANTI-epD3

Rh(D)-VARIANT VI INDIVIDUAL

Rh(D)-NEGATIVE INDIVIDUAL